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SAU 1643
IDS #4
Jina C.
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Dkt. 30634-B/JPW/JML/YL

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants : Eric Rose, et al.

Serial No. : 09/053,872

Filed : April 1, 1998 Art Unit: 1643

For : METHODS FOR INHIBITING THROMBOSIS IN A
PATIENT WHOSE BLOOD IS SUBJECTED TO
EXTRACORPOREAL CIRCULATION

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1185 Ave. of the Americas
New York, New York 10036
November 24, 1998

Assistant Commissioner for Patents
Washington, D.C. 20231

Sir:

INFORMATION DISCLOSURE STATEMENT

In accordance with their duty of disclosure under 37 C.F.R. §1.56, applicants would like to direct the Examiner's attention to the following documents, which are listed on Form PTO-1449 and are also listed below. Copies of the documents were previously submitted in connection with copending, coassigned, U.S. Serial No. 08/648,561, filed May 16, 1996, of which the above identified patent application is a continuation-in-part. The documents listed herebelow as items 1-14 are again listed on Form PTO-1449 which is attached hereto as Exhibit 1.

1. Bazan, J. F., Big rigs in blood coagulation, Nature 380:21-23, 03/1996;
2. Benedict, C.R. et al., Active site-blocked factor IXa prevents intravascular thrombus formation in the coronary vasculature without inhibiting extravascular coagulation in a canine thrombosis model, J. Clin. Invest., 88:1760-1765, 1991;

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Page 2

3. Benedict, C. R., et al., Endothelial-Dependent Procoagulant and Anticoagulant Mechanisms, Texas Heart Institute Journal, 21: 86-90, 1994;
4. Berntorp, E., Biochemical and in vivo properties of high purity factor IX concentrates, Thrombosis and Haemostasis 70(5):768-773, 11/1993;
5. Freedman, S.J. et al., Structure of the metal-free γ -carboxyglutamic acid-rich membrane binding region of factor IX by two-dimensional NMR spectroscopy, J. Biol. Chem., 270(14):7980-7987, 04/1995;
6. Furie, B.C. and Furie, B. (1995) Biosynthesis of factor IX: implications for gene therapy, Thrombosis and Haemostasis, 74(1):274-277;
7. Iberti, T.J. et al., Abnormal coagulation profile in brain tumor patients during surgery, Neurosurgery, 34(3):389-395, 03/1994;
8. Kirchhofer, et al. (1995) Active site-blocked factors VIIa and IXa differentially inhibit fibrin formation in a human ex vivo thrombosis model, Arterioscler. Thromb. Vasc. Biol., 15:1098-1106;
9. Kuwabara, K. et al., Calreticulin, an antithrombotic agent which binds to vitamin K-dependent coagulation factors, stimulates endothelial nitric oxide production, and limits thrombosis in canine coronary arteries, J. Biol. Chem., 270(14):8179-8187, 04/1995;

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Page 3

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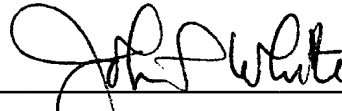
10. Miyata, T. et al., Factor IX Bm Kiryu: a Val-313-to-Asp substitution in the catalytic domain results in loss of function due to a conformational change of the surface loop: evidence obtained by chimeric modeling, Brit. J. Of Haematol., 88(1):156-165, 09/1994;
11. Santagostino, E. et al., Markers of hypercoagulability in patients with hemophilia B given repeated, large doses of factor IX concentrates during and after surgery, Thrombosis and Haemostasis, 71(6):737-40, 06/1994;
12. Wacey, A.I. et al., Determinants of the factor IX mutational spectrum in hemophilia B: an analysis of missense mutations using a multi-domain molecular model of the activated protein, Hum. Genet., 94(6):594-608, 12/1994;
13. Warriier, I. et al., Safety of high doses of a monoclonal antibody-purified factor IX concentrate, Am. J. Of Hematol., 49(1):92-94, 05/1995;
14. Wong, A. G. et al., Relative efficacy of active site-blocked factors IXa Xa in a model of venous thrombosis, Supplement I Circulation, Abstract # 3293 p. I-686, Vol. 92 No. 8, 10/1995;

Applicants request that the Examiner review the references and make them of record in the subject application.


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No fee is deemed necessary in connection with the filing of this Information Disclosure Statement. If any such fee is required, authorization is hereby given to charge the amount of such fee to Deposit Account No. 03-3125.

Respectfully submitted,



John P. White
Registration No. 28,678
Attorney for Applicants
Cooper & Dunham LLP
1185 Avenue of the Americas
New York, New York 10036
(212) 278-0400

I hereby certify that this correspondence is being deposited this date with the U.S. Postal Service with sufficient postage as first class mail in an envelope addressed to: Assistant Commissioner for Patents Washington, D.C. 20231.	
 John P. White Reg. No. 28,678	11/24/98 Date

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Exhibit 1



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Form PTO-1449 (Substituted)
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INFORMATION DISCLOSURE CITATION

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APPLICANTS
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OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

JR	Benedict, C. R., et al., Endothelial-Dependent Procoagulant and Anticoagulant Mechanisms, <u>Texas Heart Institute Journal</u> , 21: 86-90, 1994;
JR	Berntorp, E., Biochemical and in vivo properties of high purity factor IX concentrates, <u>Thrombosis and Haemostasis</u> , 70(5):768-773, 11/1993;
JR	Freedman, S.J. et al., Structure of the metal-free γ -carboxyglutamic acid-rich membrane binding region of factor IX by two-dimensional NMR spectroscopy, <u>J. Biol. Chem.</u> , 270(14):7980-7987, 04/1995;
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JR	Kirchhofer, et al. (1995) Active site-blocked factors VIIa and IXa differentially inhibit fibrin formation in a human ex vivo thrombosis model, <u>Arterioscler. Thromb. Vasc. Biol.</u> , 15:1098-1106;
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EXAMINER

Jeffrey E. Russell

DATE CONSIDERED

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Wacey, A.I. et al., Determinants of the factor IX mutational spectrum in hemophilia B: an analysis of missense mutations using a multi-domain molecular model of the activated protein, Hum. Genet., 94(6):594-608, 12/1994;

Warrier, I. et al., Safety of high doses of a monoclonal antibody-purified factor IX concentrate, Am. J. Of Hematol., 49(1):92-94, 05/1995;

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